

APPLICANT: Oxford, et al.
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Page 2

Amendments to the specification:

At page 1, line 2, insert the following:

This application is a divisional of USSN 09/964,260, filed on September 26, 2001, which in turn claimed the benefit of priority to WO 00/58308, published October 5, 2000.

At page 2, lines 1-6, amend as follows:

The potential adverse effects of a PDE III/IV inhibitor (e.g., nausea and vomiting, gastric acid secretion, cardiovascular effects such as increase cardiac contractility, vasodilation and potential arrhythmogenic activity) should be avoidable with a compound that is directly delivered to the lungs by inhalation. It is desirable that the substance is long acting[[,]] and non-irritant ~~and has a taste which is not so unpleasant as to have any adverse effect on patient compliance.~~

At page 2, lines 14-18, amend as follows:

As described by De Souza *et al.* and in GB-A-1597717, trequinsin has valuable pharmacological properties, and can be administered to human subjects suffering from, for example, respiratory disorders. However, it is unsuitable for administration by inhalation because ~~of its bitter taste and in vitro~~ data indicate its persistence of action is less than desirable.

At page 2, lines 20-23, amend as follows:

It has now been found that it is possible to design certain pyrimido [6,1-a] isoquinolin-4-one derivatives which are PDE inhibitors, which have a longer duration of action relative to trequinsin and other useful properties, ~~such as improved taste.~~

At the last page of the specification, please insert the abstract attached hereto.